L Number	Hits	Search Text	DB	Time stamp
1	2	6140446.pn.	USPAT;	2004/02/04 15:01
		1	US-PGPUB;	
			ЕРО; ЈРО;	
			DERWENT	
5	4	6242568.pn. or 6140466.pn.	USPAT;	2004/02/04 15:04
		1	US-PGPUB;	
			ЕРО; ЛРО;	
			DERWENT	
6	160	barbas-\$.in.	USPAT;	2004/02/04 15:05
			US-PGPUB;	
			ЕРО; ЈРО;	
			DERWENT	
7	63	gottesfeld-\$.in.	USPAT;	2004/02/04 15:05
			US-PGPUB;	
			ЕРО; ЈРО;	
			DERWENT	
8	814	wright-p\$.in.	USPAT;	2004/02/04 15:06
			US-PGPUB;	
			ЕРО; ЛРО;	
			DERWENT	
9	7	barbas-\$.in. and gottesfeld-\$.in. and wright-p\$.in.	USPAT;	2004/02/04 15:06
			US-PGPUB;	
:			ЕРО; ЛРО;	
			DERWENT	
10	1023	barbas-\$.in. or gottesfeld-\$.in. or wright-p\$.in.	USPAT;	2004/02/04 15:06
			US-PGPUB;	
			ЕРО; ЛРО;	
			DERWENT	
11	29		USPAT;	2004/02/04 15:06
		finger)	US-PGPUB;	
			ЕРО; ЛРО;	
			DERWENT	
12	22	((barbas-\$.in. or gottesfeld-\$.in. or wright-p\$.in.) and (zinc adj2	USPAT;	2004/02/04 15:06
		finger)) not (barbas-\$.in. and gottesfeld-\$.in. and wright-p\$.in.)	US-PGPUB;	
			ЕРО; ЈРО;	
			DERWENT	2004/02/04 15:06
13	3653	zinc adj2 finger	USPAT;	2004/02/04 15:06
			US-PGPUB;	
			EPO; JPO;	
1,,	470	(-in- adi) finan) ti	DERWENT USPAT;	2004/02/04 15:06
14	478	(zinc adj2 finger).ti.	US-PGPUB;	2004/02/04 15:00
			EPO; JPO;	
			DERWENT	
15	17	((zinc adj2 finger) with bind\$4 with (new or different)).ti.	USPAT;	2004/02/04 15:06
13	1,	((Zhio daj2 illigor) with oliday with (now or differency).	US-PGPUB;	
			EPO; JPO;	
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16	338	ladner-\$.in.	USPAT;	2004/02/04 15:07
'			US-PGPUB;	
			ЕРО; ЛРО;	
			DERWENT	
17	14	ladner-\$.in. and (zinc adj2 finger)	USPAT;	2004/02/04 15:07
		, , , ,	US-PGPUB;	
			ЕРО; ЛРО;	
			DERWENT	
18	155	(zinc adj2 finger) with librar\$4	USPAT;	2004/02/04 15:07
			US-PGPUB,	
			ЕРО; ЈРО;	
			DERWENT	

19	99	(zinc adj2 finger) with librar\$4 with (produc\$6 or screen\$4 or	USPAT;	2004/02/04 15:07
		construct\$4 or design\$4)	US-PGPUB;	
İ		,	ЕРО; ЈРО;	
			DERWENT	
20	22	((zinc adj2 finger) with librar\$4 with (produc\$6 or screen\$4 or	USPAT;	2004/02/04 15:07
		construct\$4 or design\$4)) with method	US-PGPUB;	
		<b>3</b> . , ,	ЕРО; ЈРО;	
			DERWENT	
21	18	(((zinc adj2 finger) with librar\$4 with (produc\$6 or screen\$4 or	USPAT;	2004/02/04 15:07
1		construct\$4 or design\$4)) with method) not (barbas-\$.in. or	US-PGPUB;	
		gottesfeld-\$.in. or wright-p\$.in.)	ЕРО; ЛРО;	
		\$ 1. ,	DERWENT	
22	2	((zinc adj2 finger) with librar\$4 with (produc\$6 or screen\$4 or	USPAT;	2004/02/04 15:08
	_	construct\$4 or design\$4)).ti.	US-PGPUB;	
			ЕРО; ЈРО;	
			DERWENT	
23	5	((zinc adj2 finger) with librar\$4 with (produc\$6 or screen\$4 or	USPAT;	2004/02/04 15:08
==		construct\$4 or design\$4)).clm.	US-PGPUB,	
	1	<b>3</b> * "	ЕРО; ЛРО;	
	-		DERWENT	

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09/500,700 Served Results 02/08/04

## -continued

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: Genomic DNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:61:

GCGTGGGCGG CGTGGGCG

18

- (2) INFORMATION FOR SEQ ID NO:62:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 18 base pairs
    - (B) TYPE: nucleic acid
      (C) STRANDEDNESS: single
    - (C) SIRANDEDNESSI BING
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: Genomic DNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:62:

GCGTGGGCGG GGGCGGGG

18

What is claimed is:

- 1. An isolated zinc finger-nucleotide binding polypeptide variant comprising at least three zinc finger modules that bind to a target cellular nucleotide sequence and modulate the transcriptional function of the cellular nucleotide sequence, wherein the amino acid sequence of each zinc finger module that binds a target cellular nucleotide comprises two cysteines and two histidines whereby both cysteines are amino proximal to both histidines and wherein each of three modules of said variant has at least one amino acid sequence modification.
- 2. The variant of claim 1, wherein the modulation is enhancement of transcription of a gene operatively linked to 35 the cellular nucleotide sequence.
- 3. The variant of claim 1, wherein the modulation is suppression of transcription of a gene operatively linked to the cellular nucleotide sequence.
- 4. The variant of claim 1, which is derived from a zinc 40 finger-nucleotide binding polypeptide selected from the group consisting of zif268 and TFIIIA.
- 5. The variant of claim 1, wherein the cellular nucleotide sequence is DNA.
- The variant of claim 1, wherein the cellular nucleotide 45 sequence is RNA.
- 7. The variant of claim 1, wherein the polypeptide contains a linker region between zinc fingers, the linker comprising the amino acid sequence TGEKP.
- 8. The variant of claim 1, wherein the cellular nucleotide 50 sequence is a structural gene nucleotide sequence.
- 9. The variant of claim 1, wherein the cellular nucleotide sequence is a promoter nucleotide sequence.
- 10. The variant of claim 9, wherein the promoter is an onco-promoter.
- 11. The variant of claim 10, wherein the promoter is a viral promoter.
- 12. The variant of claim 1, wherein the cellular nucleotide
- sequence is a retroviral nucleotide sequence.

  13. The variant of claim 12, wherein the retrovirus is a 60
- human T-cell lymphotrophic virus (HTLV).

  14. The variant of claim 13, wherein the retrovirus is HTLV-1 or HTLV-2.
- 15. The variant of claim 12, wherein the retrovirus is a human immunodeficiency virus (HIV).
- 16. The variant of claim 15, wherein the retrovirus is HIV-1 or HIV-2.

- 17. The variant of claim 1, wherein the cellular nucleotide sequence is an oncogene nucleotide sequence.
- 18. The variant of claim 1, wherein the cellular nucleotide sequence is a plant cellular nucleotide sequence.
- 19. The isolated zinc finger-nucleotide binding polypeptide variant of claim 1, comprising at least four zinc finger modules that bind to a cellular nucleotide sequence and modulate the transcriptional function of the cellular nucleotide sequence.

20. The isolated zinc finger-nucleotide binding polypeptide variant of claim 1, comprising at least six zinc finger modules that bind to a cellular nucleotide sequence and modulate the transcriptional function of the cellular nucleotide sequence.

- 21. The isolated zinc finger-nucleotide binding polypeptide variant of claim 1, wherein the polypeptide binds to a cellular nucleotide sequence having 18 contiguous base pairs.
- 22. The isolated zinc finger-nucleotide binding polypeptide variant of claim 1, wherein the polypeptide binds to a cellular nucleotide sequence comprising two 9-base pair binding sites.
- 23. An isolated zinc finger-nucleotide binding polypeptide variant having at least six zinc finger modules that bind to a target cellular nucleotide sequence and modulate the transcriptional function of the cellular nucleotide sequence, wherein each zinc finger module that binds a target cellular nucleotide sequence is modified, wherein the polypeptide binds to a cellular nucleotide sequence comprising two 9-base pair binding sites and wherein the two 9-base pair binding sites are separated by a variable number of nucleotides.
- 24. The isolated zinc finger-nucleotide binding polypeptide variant of claim 23, wherein the two 9-base pair binding sites are contiguous.
- 25. A n isolated nucleotide sequence encoding a zinc finger-nucleotide binding polypeptide variant of claim 1.
- 26. The nucleotide sequence of claim 19, further comprising a transcriptional activation domain in operable linkage with the nucleotide sequence.
- 27. The nucleotide sequence of claim 19, further comprising a repressor domain in operable linkage with the nucleotide sequence.
- 28. A nucleotide sequence encoding a zinc finger nucleotide binding polypeptide variant having the zinc finger

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modules that bind to a target cellular nucleotide sequence and modulate the transcriptional function of the cellular nucleotide sequence, wherein each zinc finger module that binds a target cellular nucleotide sequence is modified comprising a transcriptional activation domain in operable 5 linkage with the nucleotide sequence, wherein the transcriptional activational domain is a herpes simplex virus VP16 protein.

- 29. A nucleotide sequence encoding a zinc fingernucleotide binding polypeptide variant having zinc finger 10 modules that bind to a target cellular nucleotide sequence and modulate the transcriptional function of the cellular nucleotide sequence, wherein each zinc finger module that binds a target cellular nucleotide sequence is modified comprising a repressor domain in operable linkage with the 15 nucleotide sequence, wherein the repressor domain is the Kruppel-associated box A domain (KRAB-A).
- 30. A recombinant expression vector containing a nucleotide sequence of claim 19.
- 31. An in vitro method for inhibiting a transcriptional 20 function of a target cellular nucleotide sequence comprising a zinc finger-nucleotide binding motif, the method comprising contacting the motif with an effective amount of a zinc finger-nucleotide binding polypeptide variant comprised of at least three zinc finger molecules, wherein the amino acid sequence of each zinc finger module that binds the motif comprises two cysteines and two histidines whereby both cysteines are amino proximal to both histidines and wherein each of three modules has at least one amino acid sequence modification, thereby inhibiting a transcriptional function of 30 the sequence.
- 32. The method of claim 26, wherein the zinc finger binding polypeptide variant is a truncated zinc finger protein.
- 33. The method of claim 31, wherein the cellular nucle- 35 otide sequence is DNA.
- 34. The method of claim 31, wherein the cellular nucleotide sequence is RNA.
- 35. The method of claim 31, wherein the cellular nucleotide sequence is a structural gene nucleotide sequence.
- 36. The method of claim 31, wherein the cellular nucleotide sequence is a promoter nucleotide sequence.
- 37. The method of claim 31, wherein the cellular nucleotide sequence is an oncogene nucleotide sequence.
- 38. The method of claim 31, wherein the cellular nucleotide sequence is a plant cellular nucleotide sequence.
- 39. The method of claim 26, wherein the variant is derived from a zinc finger-nucleotide binding polypeptide selected from the group consisting of Zif268 and TFIIIA.
- 40. A method for isolating a zinc finger-nucleotide binding polypeptide variant which binds to a cellular nucleotide sequence comprising:
  - a) identifying the amino acids in a zinc finger-nucleotide binding polypeptide that bind to a first cellular nucleotide sequence and modulate the function of a nucleotide sequence;
  - b) creating an expression library encoding the polypeptide variant containing randomized substitution of the amino acids identified in step a) above;
  - c) expressing the library in a suitable host cell; and
  - d) isolating a clone that produces a polypeptide variant that binds to a second cellular nucleotide sequence and modulates the function of the second nucleotide sequence;

wherein the variant is comprised of at least three zinc finger modules and wherein the amino acid sequence of each module that binds the second nucleotide sequence comprises two cysteines and two histidines whereby both cysteines are amino proximal to both histidines and wherein each of three modules of said variant has at least one amino acid sequence modification.

- 41. The method of claim 40, wherein the library is expressed in a phage surface expression system.
- 42. The method of claim 41, wherein the phage expression system includes a reducing reagent which allows folding of expression products on the phage surface.
- 43. The method of claim 42, wherein the reducing reagent is dithiothreitol.
- 44. The method of claim 40, wherein the library is randomized by PCR using primers containing degenerate triplet codons at sequence locations corresponding to the determined amino acids.
- 45. The method of claim 40, wherein the variant is derived from a zinc finger-nucleotide binding polypeptide selected from the group consisting of zif268 and TFIIIA.
- 46. The method of claim 45, wherein the variant derived from the zinc finger-nucleotide binding polypeptide zif268 is modified at any of residues 1, 3, 4, 5, 6 or 7 as set forth in SEQ ID NO:14.
- 47. The method of claim 45, wherein the variant derived from the zinc finger-nucleotide binding polypeptide zif268 is modified at any of residues 1, 2, 3, 4, 5 or 6 as set forth in SEQ ID NO:15.
- 48. The method of claim 45, wherein the variant derived from the zinc finger-nucleotide binding polypeptide zif268 is modified at any of residues 1, 3, 4, 5, 6 or 7 as set forth SEQ ID NO:5.
  - 49. The method of claim 40, wherein the modulation of function is enhancement of transcription of a gene operatively linked to the cellular nucleotide sequence.
  - 50. The method of claim 40, wherein the modulation of function is suppression of transcription of a gene operatively linked to the cellular nucleotide sequence.
  - 51. The method of claim 40, wherein the cellular nucle-40 otide sequence is DNA.
    - 52. The method of claim 40, wherein the cellular nucleotide sequence is RNA.
    - 53. A zinc finger-nucleotide binding polypeptide variant produced by the method of claim 40.
    - 54. A method for identifying a zinc finger-nucleotide binding polypeptide variant comprised of at least three zinc finger modules, which modulates the transcriptional function of cellular nucleotide sequence and binds to a zinc finger-nucleotide binding motif, wherein the amino acid sequence of each module that binds to a zinc finger nucleotide binding motif comprises two cysteines and two histidines whereby both cysteines are amino proximal to both histidines and wherein each of three modules of said variant has at least one amino acid sequence modification, said method comprising:
      - a) incubating the components comprising a nucleotide sequence encoding the putative modulating variant operably linked to a first inducible promoter, and a reporter gene operably linked to a second inducible promoter and a zinc finger-nucleotide binding motif, wherein the incubating is carried out under conditions sufficient to allow the components to interact; and
      - b) measuring the effect of the putative modulating variant on the expression of the reporter gene.

\* \* \* \* \*

## US 6,242,568 B1

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Arg	Asn 130	Phe	Ser	Arg	Ser	<b>Asp</b> 135	His	Leu	Thr	Thr	Нів 140	Ile	Arg	Thr	His	
ACA	GGC	GAG	AAG	CCT	TTT	GCC	TGT	GAC	ATT	TGT	GGG	AGG	AAG	TTT	GCC	480
Thr	Gly	Glu	Lys	Pro	Phe	Ala	Сув	Αsp	Ile	Сув	Gly	Arg	Lys	Phe	Ala	
145					150					155					160	
AGG	AGT	GAT	GAA	CGC	AAG	AGG	CAT	ACC	AAA	ATC	CAT	TTA	AGA	CAG	AAG	528
Arg	Ser	qaA	Glu	-	Lys	Arg	Нiв	Thr		Ile	His	Leu	Arg		Ļув	
				165					170					175		
GAC	TCT	AGA	ACT	AGT												543
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			20					25					30			
Gln	Lys	Pro	Phe	Gln	Сув	Arg	Ile	Сув	Met	Arg	Asn	Phe	Ser	Arg	Ser	
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Aen	His	Len	Thr	Thr	нів	Ile	Ara	Thr	His	Thr	Glv	Glu	Lvs	Pro	Phe	
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Ala 65	Cys	Авр	Ile	Сув	G1y 70	Arg	ГÀВ	Phe	ALA	Arg 75	Ser	Авр	GIU	Arg	80 FÅR	
Arg	His	Thr	Lys		His	Thr	Gly	Glu	aęı 90	Pro	Tyr	Ala	Сув	Pro 95	Val	
				85					90					,,		
Glu	Ser	Сув	Asp	Arg	Arg	Phe	Ser	Arg	Ser	Авр	Glu	Leu	Thr	Arg	His	
			100					105					110			
Ile	Arg	Ile	His	Thr	Gly	Gln	Lув	Pro	Phe	Gln	Сув	Arg	Ile	Cys	Met	
	•	115			-		120					125				
Ara	Asn	Dhe	Ser	Arc	Sar	Asp	Hie	Len	Thr	Thr	Hie	Ile	Ara	Thr	His	
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Thr 145	Gly	GIu	Lys	Pro	Phe 150	ATG	Сув	Авр	TIE	Cys 155	GTÀ	Arg	гув	rne	160	
Arg	Ser	Asp	Glu		Lys	Arg	His	Thr	Lys 170	Ile	His	Leu	Arg	Gln 175	Lys	
				165					110					-,,		
Авр	Ser	Arg	Thr	Ser												

What is claimed is:

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1. A method for isolating a zinc finger-nucleotide binding polypeptide variant which binds to a cellular nucleotide sequence comprising:

- a) identifying the amino acids in a zinc finger-nucleotide binding polypeptide that bind to a first cellular nucleotide sequence and modulate the function of a nucleotide sequence;
- b) creating an expression library encoding the polypeptide variant containing randomized substitution of the 65 amino acids identified in step a) above;
- c) expressing the library in a suitable host cell; and

- d) isolating a clone that produces a polypeptide variant that binds to a second cellular nucleotide sequence and modulates the function of the second nucleotide sequence:
- wherein the variant is comprised of at least two zinc finger modules and wherein the amino acid sequence of each module that binds the second nucleotide sequence comprises two cysteines and two histidines whereby both cysteines are amino proximal to both histidines and wherein each of two modules of said variant has at least one amino acid sequence modification.
- 2. The method of claim 1, wherein the library is expressed in a phage surface expression system.

- 3. The method of claim 1, wherein the phage expression system includes a reducing reagent which allows folding of expression products on the phage surface.
- 4. The method of claim 3, wherein the reducing reagent is dithiothreitol.
- 5. The method of claim 1, wherein the library is randomized by PCR using primers containing degenerate triplet codons at sequence locations corresponding to the determined amino acids.
- 6. The method of claim 1, wherein the modulation of function is enhancement of transcription of a gene operatively linked to the cellular nucleotide sequence
- 7. The method of claim 1, wherein the modulation of function is suppression of transcription of a gene operatively linked to the cellular nucleotide sequence.
- sequence is DNA.
- 9. The method of claim 1, wherein the cellular nucleotide sequence is RNA.
- 10. The method of claim 1, wherein the variant is derived from a zinc finger-nucleotide binding polypeptide selected 20 from the group consisting of zif 268 and TFIIIA.
- 11. The method of claim 1, wherein the variant derived from the zinc finger-nucleotide binding polypeptide zif268 is modified at any of residues 1, 3, 4, 5, 6 or 7 as set forth in SEQ ID NO:14.
- 12. The method of claim 1, wherein the variant derived from the zinc finger-nucleotide binding polypeptide zif268 is modified at any of residues 1, 2, 3, 4, 5 or 6 as set forth in SEQ ID NO:15.
- 13. The method of claim 1, wherein the variant derived 30 from the zinc finger-nucleotide binding polypeptide zif268 is modified at any of residues 1, 3, 4, 5, 6 or 7 as set forth in SEQ ID NO:5.
- 14. A method for identifying a zinc finger-nucleotide binding polypeptide variant comprised of at least two zinc finger modules, which modulates the transcriptional function of cellular nucleotide sequence and binds to a zinc finger-nucleotide binding motif, wherein the amino acid sequence of each module that binds to a zinc finger nucleotide binding motif comprises two cysteines and two histidines whereby both cysteines are amino proximal to both histidines and wherein each of two modules of said variant has at least one amino acid sequence modification, said method comprising:
  - a) incubating the components comprising a nucleotide 45 sequence encoding the putative modulating variant operably linked to a first inducible promoter, and a reporter gene operably linked to a second inducible promoter and a zinc finger-nucleotide binding motif, wherein the incubating is carried out under conditions 50 otide sequence is RNA. sufficient to allow the components to interact; and
  - b) measuring the effect of the putative modulating variant on the expression of the reporter gene.
- 15. The method of claim 14, wherein the modulation is inhibition of gene expression.
- 16. The method of claim 14, wherein the modulation is enhancement of gene expression.
- 17. The method of claim 14, wherein the first inducible promoter is the arabinase promoter.
- 18. The method of claim 14, wherein the second inducible 60
- promoter is the lactose promoter. 19. The method of claim 14, wherein the incubating is performed in vitro.
- 20. The method of claim 14, wherein the incubating is performed in vivo.
- 21. The method of claim 14, wherein the reporter gene is β-galactosidase.

- 22. A method of modulating transcription of a cellular nucleotide sequence associated with a zinc finger-nucleotide binding motif, comprising contacting the zinc fingernucleotide in cells in vitro with an effective amount of a zinc finger-nucleotide binding polypeptide variant comprised of at least two zinc finger modules that binds to the zinc finger-nucleotide binding motif, wherein the amino acid sequence of each module that binds the motif comprises two cysteines and two histidines whereby both cysteines are amino proximal to both histidines and wherein each of two modules of said variant has at least one amino acid sequence modification, thereby modulating transcriptional activity of the cellular nucleotide sequence.
- 23. The method of claim 22, wherein an expression vector comprising a polynucleotide sequence encoding a zinc 8. The method of claim 1, wherein the cellular nucleotide 15 finger-nucleotide binding polypeptide variant is introduced into the cells.
  - 24. The method of claim 23, wherein the expression vector is a virus.
  - 25. The method of claim 22, wherein the modulation is enhancement of transcription of a gene operatively linked to the cellular nucleotide sequence.
  - 26. The method of claim 22, wherein the modulation is suppression of transcription of a gene operatively linked to the cellular nucleotide sequence.
  - 27. The method of claims 1, 14, or 22 wherein the variant is derived from a zinc finger-nucleotide binding polypeptide selected from the group consisting of zif268 and TFIIIA.
  - 28. An isolated zinc finger-nucleotide binding polypeptide variant comprising at least two zinc finger modules that bind to a target cellular nucleotide sequence and modulate the transcriptional function of the cellular nucleotide sequence, wherein the amino acid sequence of each zinc finger module that binds a target cellular nucleotide comprises two cysteines and two histidines whereby both cysteines are amino proximal to both histidines and wherein each of two modules of said variant has at least one amino acid sequence modification.
  - 29. The variant of claim 28, wherein the modulation is enhancement of transcription of a gene operatively linked to 40 the cellular nucleotide sequence.
    - 30. The variant of claim 28, wherein the modulation is suppression of transcription of a gene operatively linked to the cellular nucleotide sequence.
    - 31. The variant of claim 28, which is derived from a zinc finger-nucleotide binding polypeptide selected from the group consisting of zif 268 and TFIIIA.
    - 32. The variant of claim 28, wherein the cellular nucleotide sequence is DNA.
    - 33. The variant of claim 28, wherein the cellular nucle-
    - 34. The variant of claim 1, wherein the polypeptide contains a linker region between zinc fingers comprising the amino acid sequence TGEKP.
  - 35. The variant of claim 28, wherein the cellular nucle-55 otide sequence is a structural gene nucleotide sequence.
    - 36. The variant of claim 28, wherein the cellular nucleotide sequence is a promoter nucleotide sequence.
    - 37. The variant of claim 36, wherein the promoter is an onco-promoter.
    - 38. The variant of claim 37, wherein the promoter is a viral promoter.
    - 39. The variant of claim 28, wherein the cellular nucleotide sequence is a retroviral nucleotide sequence.
  - 40. The variant of claim 39, wherein the retrovirus is a 65 human T-cell lymphotrophic virus (HTLV).
    - 41. The variant of claim 40, wherein the retrovirus is HTLV-1 or HTLV-2.

- 42. The variant of claim 39, wherein the retrovirus is a human immunodeficiency virus (HIV).
- 43. The variant of claim 42, wherein the retrovirus is HIV-1 or HIV-2.
- 44. The variant of claim 28, wherein the cellular nucle- 5 otide sequence is an oncogene nucleotide sequence.
- 45. The variant of claim 28, wherein the cellular nucleotide sequence is a plant cellular nucleotide sequence.
- 46. A zinc finger-nucleotide binding polypeptide variant produced by the method of claim 36.
- 47. An isolated nucleotide sequence encoding a zinc finger-nucleotide binding polypeptide variant of claim 28.
- 48. A recombinant expression vector containing a nucleotide sequence of claim 47.
- 49. A method for inhibiting a transcriptional function of a 15 otide sequence is a promoter nucleotide sequence. target cellular nucleotide sequence comprising a zinc fingernucleotide binding motif, the method comprising contacting the motif with an effective amount of a zinc fingernucleotide binding polypeptide variant comprised of at least two zinc finger molecules, wherein the amino acid sequence 20 of each zinc finger module that binds the motif comprises

two cysteines and two histidines whereby both cysteines are amino proximal to both histidines and wherein each of two modules has at least one amino acid sequence modification.

- 50. The method of claim 26, wherein the zinc finger binding polypeptide variant is a truncated zinc finger protein.
- 51. The method of claim 49, wherein the cellular nucleotide sequence is DNA.
- 52. The method of claim 49, wherein the cellular nucleotide sequence is RNA.
- 53. The method of claim 49, wherein the cellular nucleotide sequence is a structural gene nucleotide sequence.
- 54. The method of claim 49, wherein the cellular nucle-
- 55. The method of claim 49, wherein the cellular nucleotide sequence is an oncogene nucleotide sequence.
- 56. The method of claim 49, wherein the cellular nucleotide sequence is a plant cellular nucleotide sequence.

02/04/2004, EAST Version: 1.4.1

## (FILE 'HOME' ENTERED AT 17:08:19 ON 08 FEB 2004)

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FILE 'MEDLINE, EMBASE, BIOSIS, CAPLUS' ENTERED AT 17:09:06 ON 08 FEB 2004
          1500 S (BARBAS, ?)/IN,AU
L1
           987 S (GOTTESFELD, ?)/IN,AU
L2
         95284 S (WRIGHT, ?)/IN,AU
L3
            5 S L1 AND L2 AND L3
L4
             4 DUPLICATE REMOVE L4 (1 DUPLICATE REMOVED)
L5
L6
         97715 S L1 OR L2 OR L3
L7
            0 S ZINC ADJ2 FINGER
L8
         26640 S ZINC (2W) FINGER
L9
           276 S L8 AND L6
            19 S L9 AND ZIF268
L10
             8 DUPLICATE REMOVE L10 (11 DUPLICATES REMOVED)
L11
             5 S L11 NOT L4
L12
L13
        . 150 S TFIIIA AND L6
             4 S L13 AND (MUTANT OR VARIANT)
L14
             4 S L14 NOT (L4 OR L10)
L15
             1 DUPLICATE REMOVE L14 (3 DUPLICATES REMOVED)
L16
L17
          1416 S L8 (S) (MUTANT OR VARIANT)
L18
           431 S L17 AND PY<1996
            17 S L18 AND TFIIIA
L19
            8 DUPLICATE REMOVE L19 (9 DUPLICATES REMOVED)
L20
            8 S L20 NOT (LL4 OR L10 OR L16)
L21
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